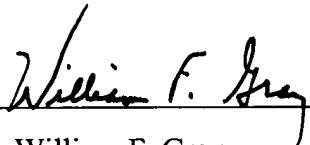


The amendment to structure (A1) in claim 1 is to clarify the drawing. Swiss-type use claims 1-8 have been converted into method of treatment claims. Claims 9-11 now refer to pharmaceutical compositions rather than medicaments. Claim 12 now refers to a process for preparing the pharmaceutical composition.

Attached hereto is a marked-up version of the changes made to the specification and claims by the current amendment. The attached page is captioned "**Version with markings to show changes made.**"

In view of the above amendments and explanations, this application is deemed to be in condition for allowance, and allowance is accordingly requested.

Respectfully submitted,



William F. Gray

Reg. No. 31018

Phone: (203) 812-2712

Date: 20 Dec '01

Bayer Corporation

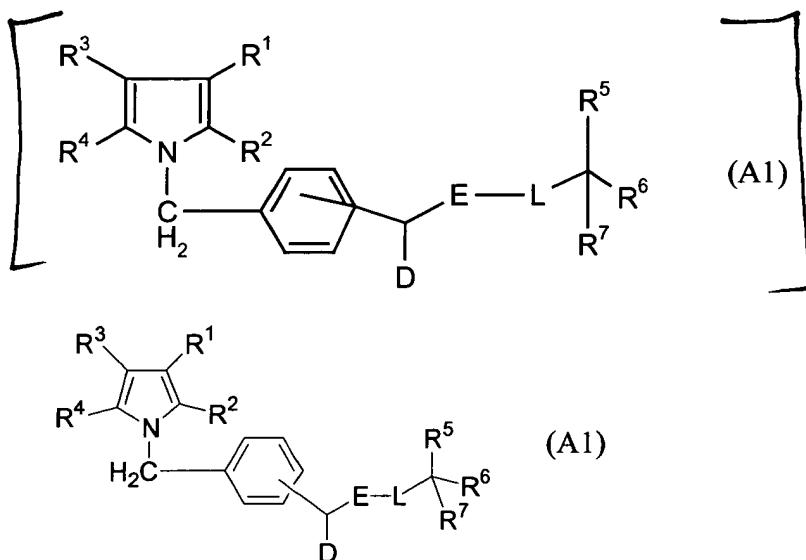
400 Morgan Lane

West Haven, CT 06516-4175

Version with markings to show changes made:In the claims:

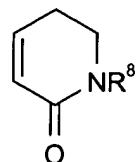
Claims 1-12 have been amended as shown below:

1. (Amended) [Use of] A method for the prophylaxis or treatment of cardiovascular diseases which comprises administering an effective amount of a combination of component A and component B, component A being at least one MTP inhibitor [as component A] of the general formula (A1)



in which

R^1 and R^2 , including the double bond connecting them, together form a phenyl or pyridyl ring or a ring of the formula



in which

R⁸ denotes hydrogen or straight-chain or branched alkyl having up to 4 carbon atoms,

R³ and R⁴, including the double bond connecting them, together form a phenyl ring or a 4- to 8-membered cycloalkene or oxocycloalkene radical,

all ring systems mentioned under R¹/R² and R³/R⁴ optionally being substituted up to 3 times, identically or differently, by halogen, trifluoromethyl, carboxyl, hydroxyl, by straight-chain or branched alkoxy or alkoxy carbonyl each having up to 6 carbon atoms or by straight-chain or branched alkyl having up to 6 carbon atoms, which for its part can be substituted by hydroxyl or by straight-chain or branched alkoxy having up to 4 carbon atoms,

D represents hydrogen, cycloalkyl having 4 to 12 carbon atoms or straight-chain or branched alkyl having up to 12 carbon atoms,

E represents the -CO- or -CS- group,

L represents an oxygen or sulphur atom or a group of the formula -NR⁹,

in which

R⁹ denotes hydrogen or straight-chain or branched alkyl having up to 6 carbon atoms, which is optionally substituted by hydroxyl or phenyl,

R⁵ denotes phenyl or a 5- to 7-membered saturated or unsaturated heterocycle having up to 3 heteroatoms from the group consisting of S, N and/or O,

the cyclic systems optionally being substituted up to 3 times, identically or differently, by nitro, carboxyl, halogen, cyano or by straight-chain or branched alkenyl or alkoxycarbonyl each having up to 6 carbon atoms or by straight-chain or branched alkyl having up to 6 carbon atoms, which is optionally substituted by hydroxyl, carboxyl or by straight-chain or branched alkoxy or alkoxycarbonyl each having up to 6 carbon atoms,

and/or the cyclic systems optionally being substituted by a group of the formula -OR¹⁰ or -NR¹¹R¹²,

in which

R¹⁰ denotes hydrogen or straight-chain or branched alkyl or alkenyl each having up to 6 carbon atoms,

R¹¹ and R¹² are identical or different and denote phenyl, hydrogen or straight-chain or branched alkyl having up to 6 carbon atoms or straight-chain or branched acyl having up to 8 carbon atoms, which is optionally substituted by a group of the formula -NR¹³R¹⁴,

in which

R¹³ and R¹⁴ are identical or different and denote hydrogen or straight-chain or branched acyl having up to 8 carbon atoms,

R⁶ represents hydrogen, carboxyl or straight-chain or branched alkoxycarbonyl having up to 5 carbon atoms,

or represents straight-chain or branched alkyl having up to 6 carbon atoms, which is optionally substituted by hydroxyl or by a group of the formula -O-CO-R¹⁵,

in which

R¹⁵ denotes phenyl which is optionally substituted up to 3 times, identically or differently, by halogen, hydroxyl or by straight-chain or branched alkyl having up to 5 carbon atoms,
 or denotes straight-chain or branched alkyl or alkenyl each having up to 22 carbon atoms, each of which is optionally substituted by a group of the formula -OR¹⁶,

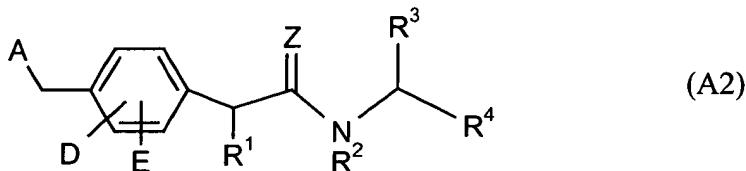
in which

R¹⁶ denotes hydrogen, benzyl, triphenylmethyl or straight-chain or branched acyl having up to 6 carbon atoms,

R⁷ represents hydrogen or

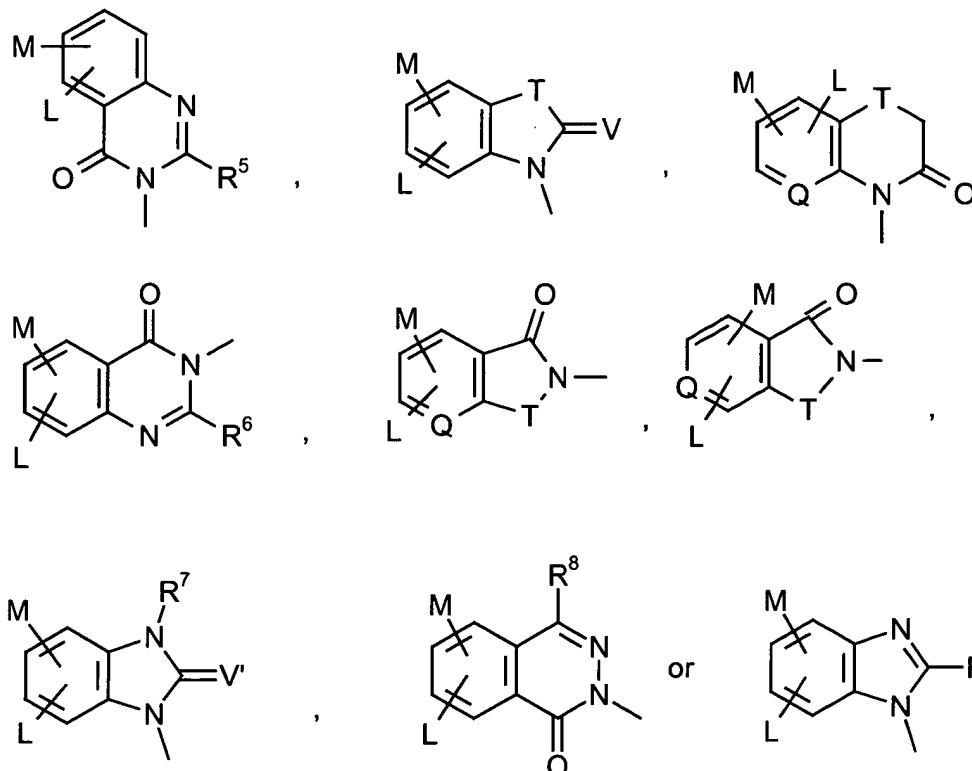
R⁶ and R⁷ together represent the group of the formula =O,

or of the general formula (A2)



in which

A represents a radical of the formula



in which

L and M are identical or different and

denote hydrogen, halogen, trifluoromethyl, carboxyl, cycloalkyl having 3 to 6 carbon atoms, hydroxyl, phenyl or straight-chain or branched alkyl, alkoxy carbonyl or alkoxy each having up to 6 carbon atoms,

Q denotes a nitrogen atom or the -CH- group,

T denotes a group of the formula -SO₂ or -CO or an oxygen or sulphur atom,

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V denotes an oxygen or sulphur atom,

R⁵, R⁶, R⁷ and R⁸ are identical or different and denote hydrogen, or straight-chain or branched alkyl having up to 6 carbon atoms, benzyl or phenyl, each of which is optionally substituted by halogen or by straight-chain or branched alkyl having up to 6 carbon atoms,

R⁹ denotes trifluoromethyl, benzyl or a 5- to 7-membered, optionally benzofused heterocycle having up to 3 heteroatoms from the group consisting of S, N and/or O, which is optionally substituted up to 3 times, identically or differently, by halogen, phenyl, hydroxyl or by straight-chain or branched alkyl or alkoxy each having up to 4 carbon atoms, or denotes a group of the formula -S(O)_a-R¹⁰,

in which

a denotes a number 0, 1 or 2,

R¹⁰ denotes straight-chain or branched alkyl or alkenyl each having up to 8 carbon atoms, each of which is optionally substituted by straight-chain or branched acyl having up to 6 carbon atoms or by aryl or aroyl each having up to 10 carbon atoms, which for their part can be substituted up to 2 times, identically or differently, by halogen, trifluoromethyl or by straight-chain or branched acyl having up to 5 carbon atoms, or denotes aryl having 6 to 10 carbon atoms, which is optionally substituted by halogen, hydroxyl, trifluoromethyl or straight-chain

or branched alkyl or alkoxy each having up to 5 carbon atoms,

D and E are identical or different and

represent hydrogen, halogen, trifluoromethyl, hydroxyl, carboxyl or straight-chain or branched alkyl, alkoxy or alkoxy carbonyl each having up to 6 carbon atoms,

Z represents an oxygen or sulphur atom,

R¹ represents cycloalkyl having 3 to 10 carbon atoms or straight-chain or branched alkyl having 1 to 10 carbon atoms, or represents phenyl which is optionally substituted up to 2 times, identically or differently, by halogen, nitro, cyano, hydroxyl, straight-chain or branched alkyl or alkoxy each having up to 4 carbon atoms,

R² represents hydrogen or straight-chain or branched alkyl having up to 3 carbon atoms,

R³ represents hydrogen or straight-chain or branched alkyl having up to 5 carbon atoms, or represents cycloalkyl having 3 to 7 carbon atoms, or represents phenyl or a 5- to 7-membered aromatic heterocycle having up to 3 heteroatoms from the group consisting of S, N and/or O, each of which is optionally substituted up to 3 times, identically or differently, by halogen, nitro, phenyl, hydroxyl or by straight-chain or branched alkyl or alkoxy having up to 6 carbon atoms,

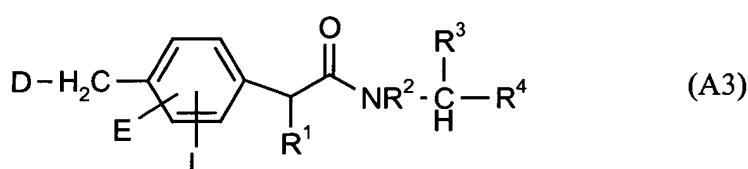
R⁴ represents hydrogen or a group of the formula -CH₂-OH or CH₂O-CO-R¹¹,

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in which

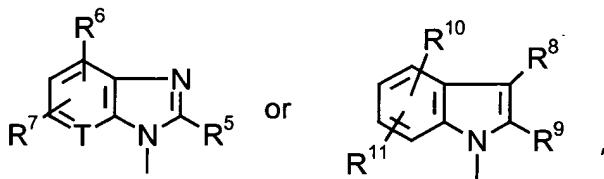
R^{11} denotes hydrogen, straight-chain or branched alkyl having up to 8 carbon atoms or phenyl which is optionally substituted up to 3 times, identically or differently, by halogen, hydroxyl, cyano or straight-chain or branched alkyl or alkoxy each having up to 4 carbon atoms,

or of the general formula (A3)



in which

D represents a radical of the formula



in which

T denotes a nitrogen atom or the -CH- group,

R^6 , R^7 , R^{10} and R^{11} are identical or different and

denote hydrogen, trifluoromethyl, halogen or straight-chain or branched

alkyl or alkoxy each having up to 6 carbon atoms,

R^5 , R^8 and R^9 are identical or different and

denote hydrogen, cycloalkyl having 3 to 6 carbon atoms, phenyl, straight-chain or branched alkoxy carbonyl having up to 6 carbon atoms or straight-chain or branched alkyl having up to 6 carbon atoms, which is optionally substituted by halogen,

or, if T represents a nitrogen atom, R^5 can also denote benzyl,

E and L are identical or different and

represent hydrogen, halogen, trifluoromethyl, hydroxyl, carboxyl or straight-chain or branched alkyl, alkoxy or alkoxy carbonyl each having up to 6 carbon atoms,

R^1 represents cycloalkyl having 3 to 10 carbon atoms or straight-chain or branched alkyl having 1 to 10 carbon atoms, or represents phenyl which is optionally substituted up to 2 times, identically or differently, by halogen, cyano, hydroxyl, straight-chain or branched alkyl or alkoxy each having up to 4 carbon atoms,

R^2 represents hydrogen or straight-chain or branched alkyl having up to 3 carbon atoms,

R^3 represents hydrogen or straight-chain or branched alkyl having up to 5 carbon atoms, or represents cycloalkyl having 3 to 7 carbon atoms, or represents phenyl or a 5- to 7-membered aromatic heterocycle having up to

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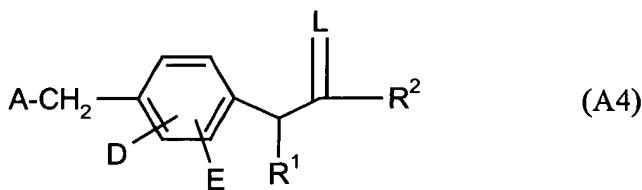
3 heteroatoms from the group consisting of S, N and/or O, each of which is optionally substituted up to 3 times, identically or differently, by halogen, nitro, phenyl, hydroxyl or by straight-chain or branched alkyl or alkoxy having up to 6 carbon atoms,

R⁴ represents hydrogen or a group of the formula -CH₂-OH or CH₂O-CO-R¹²,

in which

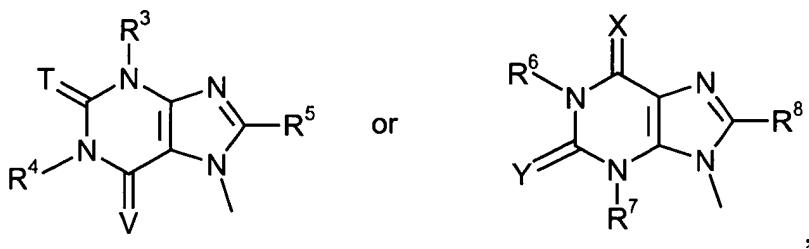
R¹² denotes hydrogen, straight-chain or branched alkyl having up to 8 carbon atoms or phenyl which is optionally substituted up to 3 times, identically or differently, by halogen, hydroxyl, cyano or straight-chain or branched alkyl or alkoxy each having up to 4 carbon atoms,

or of the general formula (A4)



in which

A represents a radical of the formula



in which

R^3 , R^4 , R^6 and R^7 are identical or different and

denote hydrogen, cycloalkyl having 3 to 7 carbon atoms or aryl having 6 to 10 carbon atoms,

or denote straight-chain or branched alkyl or alkenyl each having up to 8 carbon atoms, each of which is optionally substituted by halogen, hydroxyl or aryl having 6 to 10 carbon atoms,

T , V , X and Y are identical or different and denote an oxygen or sulphur atom,

R^5 and R^8 are identical or different and

denote hydrogen, halogen, cycloalkyl having 3 to 8 carbon atoms or straight-chain or branched alkyl or alkenyl each having up to 8 carbon atoms, each of which is optionally substituted by cycloalkyl having 3 to 8 carbon atoms, or by a 5- to 6-membered, aromatic, optionally benzofused heterocycle having up to 3 heteroatoms from the group consisting of S, N and/or O, or by aryl having 6 to 10 carbon atoms, where the cyclic systems for their part can be substituted up to 3 times, identically or differently, by a 5- to 6-membered aromatic heterocycle having up to 3 heteroatoms from the group consisting of S, N and/or O, or by phenyl, benzyl, halogen, hydroxyl, carboxyl or by straight-chain or branched alkyl, alkoxy or alkoxy carbonyl each having up to 6 carbon atoms, or

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denote aryl having 6 to 10 carbon atoms or a 5- to 7-membered aromatic, optionally benzo-fused heterocycle having up to 3 heteroatoms from the group consisting of S, N and/or O, each of which is optionally substituted up to 3 times, identically or differently, by halogen, phenyl, trifluoromethyl, hydroxyl, carboxyl or by straight-chain or branched alkyl, alkoxy or alkoxycarbonyl each having up to 6 carbon atoms or by a group of the formula -(CO)_a-NR⁹R¹⁰,

in which

a denotes a number 0 or 1,

R⁹ and R¹⁰ are identical or different and

denote hydrogen, phenyl or straight-chain or branched alkyl or acyl each having up to 5 carbon atoms,

D and E are identical or different and

represent hydrogen, halogen, trifluoromethyl, hydroxyl, carboxyl or straight-chain or branched alkyl, alkoxy or alkoxycarbonyl each having up to 6 carbon atoms,

R¹ represents hydrogen or cycloalkyl having 3 to 8 carbon atoms, or represents straight-chain or branched alkyl or alkenyl each having up to 8 carbon atoms, each of which is optionally substituted by cycloalkyl having 3 to 6 carbon atoms, phenyl or by a 5- to 6-membered aromatic heterocycle having up to 3 heteroatoms from the group consisting of S, N and/or O, or represents phenyl or a 5- to 6-membered aromatic heterocycle having up to 3 heteroatoms from the group consisting of S, N and/or O, the ring systems optionally being substituted up to 3 times, identically or differently, by halogen,

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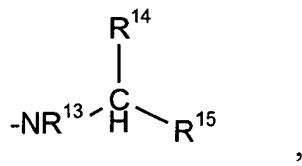
phenyl, trifluoromethyl or straight-chain or branched alkyl or alkoxy each having up to 5 carbon atoms, hydroxyl or by a group of the formula -NR¹¹R¹²,

in which

R¹¹ and R¹² have the meaning of R⁹ and R¹⁰ indicated above and are identical to or different from this,

L represents an oxygen or sulphur atom,

R² represents mercapto, hydroxyl, straight-chain or branched alkoxy having up to 8 carbon atoms or the group of the formula



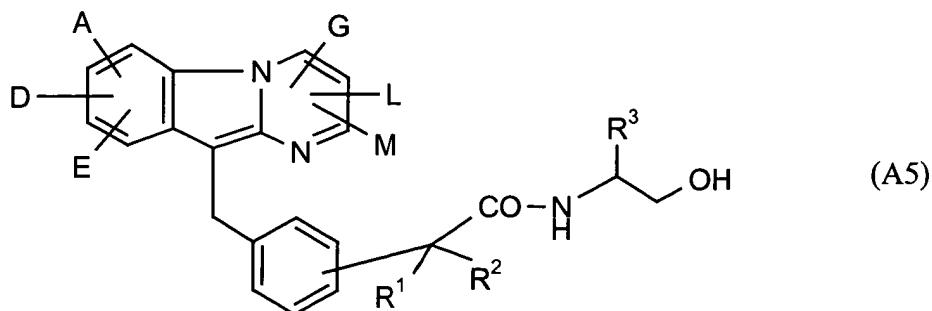
in which

R¹³ denotes hydrogen or straight-chain or branched alkyl having up to 4 carbon atoms,

R¹⁴ denotes hydrogen, phenyl or a 5- to 6-membered aromatic heterocycle having up to 3 heteroatoms from the group consisting of S, N and/or O,

R¹⁵ denotes hydrogen or straight-chain or branched alkyl having up to 8 carbon atoms, which is optionally substituted by hydroxyl,

or of the general formula (A5)



in which

A, D, E, G, L and M are identical or different and

represent hydrogen, halogen, trifluoromethyl, carboxyl, hydroxyl, straight-chain or branched alkoxy or alkoxy carbonyl each having up to 6 carbon atoms or straight-chain or branched alkyl having up to 6 carbon atoms, which for its part can be substituted by hydroxyl or by straight-chain or branched alkoxy having up to 4 carbon atoms,

R¹ and R² are identical or different and

represent hydrogen, cycloalkyl having 3 to 8 carbon atoms or straight-chain or branched alkyl having up to 10 carbon atoms, which is optionally substituted by cycloalkyl having 3 to 6 carbon atoms or represent phenyl which is optionally substituted by halogen or trifluoromethyl, or

R¹ and R², together with the carbon atom, form a 4- to 8-membered cycloalkyl ring

and

R³ represents phenyl which is optionally substituted up to 3 times, identically or differently, by nitro, carboxyl, halogen, cyano or by straight-chain or branched alkenyl or alkoxycarbonyl each having up to 6 carbon atoms or by straight-chain or branched alkyl having up to 6 carbon atoms, which is optionally substituted by hydroxyl, carboxyl or by straight-chain or branched alkoxy or alkoxycarbonyl each having up to 6 carbon atoms,
and/or is optionally substituted by a group of the formula -OR⁴ or -NR⁵R⁶,

in which

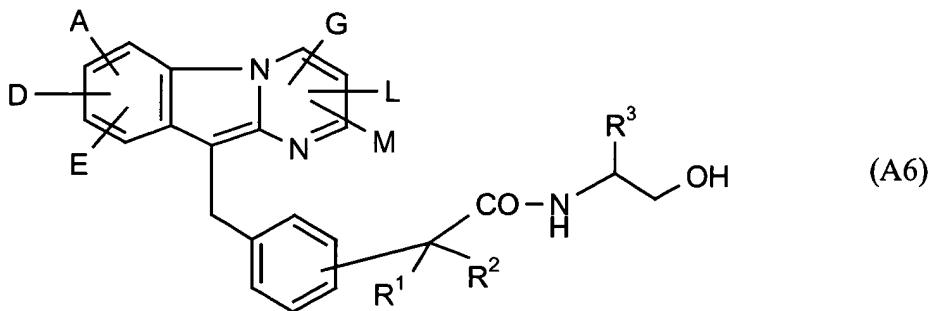
R⁴ denotes hydrogen or straight-chain or branched alkyl or alkenyl each having up to 6 carbon atoms,

R⁵ and R⁶ are identical or different and denote phenyl, hydrogen or straight-chain or branched alkyl having up to 6 carbon atoms,
or denote straight-chain or branched acyl having up to 8 carbon atoms,
which is optionally substituted by a group of the formula -NR⁷R⁸,

in which

R⁷ and R⁸ are identical or different and
denote hydrogen or straight-chain or branched acyl having up to 8 carbon atoms,

or of the general formula (A6)



in which

A, D, E, G, L and M are identical or different and

represent hydrogen, halogen, trifluoromethyl, carboxyl, hydroxyl, straight-chain or branched alkoxy or alkoxycarbonyl each having up to 6 carbon atoms or straight-chain or branched alkyl having up to 6 carbon atoms, which for its part can be substituted by hydroxyl or by straight-chain or branched alkoxy having up to 4 carbon atoms,

R¹ and R² are identical or different and

represent hydrogen, cycloalkyl having 3 to 8 carbon atoms or straight-chain or branched alkyl having up to 10 carbon atoms, which is optionally substituted by cycloalkyl having 3 to 6 carbon atoms, or
 represent phenyl which is optionally substituted by halogen or trifluoromethyl, or

R¹ and R², together with the carbon atom, form a 4- to 8-membered cycloalkyl ring

and

R³ represents phenyl which is optionally substituted up to 3 times, identically or differently, by nitro, carboxyl, halogen, cyano or by straight-chain or branched alkenyl or alkoxycarbonyl each having up to 6 carbon atoms or by straight-chain

or branched alkyl having up to 6 carbon atoms, which is optionally substituted by hydroxyl, carboxyl or by straight-chain or branched alkoxy or alkoxycarbonyl each having up to 6 carbon atoms,

and/or is optionally substituted by a group of the formula -OR⁴ or -NR⁵R⁶,

in which

R⁴ denotes hydrogen or straight-chain or branched alkyl or alkenyl each having up to 6 carbon atoms,

R⁵ and R⁶ are identical or different and denote phenyl, hydrogen or straight-chain or branched alkyl having up to 6 carbon atoms,
or denote straight-chain or branched acyl having up to 8 carbon atoms,
which is optionally substituted by a group of the formula -NR⁷R⁸,

in which

R⁷ and R⁸ are identical or different and
denote hydrogen or straight-chain or branched acyl having up to 8 carbon atoms,

if appropriate in an isomeric form and their salts

and component B being at least one [with] HMG-CoA reductase inhibitor[s] [as component B for the production of medicaments for the prophylaxis and/or treatment of cardiovascular diseases].

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2. (Amended) [Use of a combination] The method according to Claim 1 [for the production of medicaments for the control or prophylaxis of] wherein said cardiovascular diseases [which] are associated with metabolic diseases or deficits.
 3. (Amended) [Use of a combination] The method according to Claim 2 for the control of arteriosclerosis, diseases of the coronary vessels of the heart, raised serum lipids, hypercholesterolaemia, hypertriglyceridaemia and mixed forms which are combined with raised VLDL or LDL and/or raised chylomicrons, and of syndrome X.
 4. (Amended) [Use of a combination] The method according to Claim 2 for the treatment of secondary hypercholesterolaemia and secondary hypertriglyceridaemia, which are optionally associated with apolipoprotein E polymorphism, obesity, chylomicronaemia and chylomicronaemia syndrome, renal insufficiency, chronic renal insufficiency, nephrotic syndrome, diabetes mellitus type II, and with hepatomas and plasmacytomas.
 5. (Amended) [Use of a combination] The method according to Claim 2, characterized in that [it contains, as] component A[,] is a compound of the general formula (A1).
 6. (Amended) [Use of a combination] The method according to Claim 2, characterized in that [it contains, as] component A[,] is a compound of Examples 1-119.
 7. (Amended) [Use of a combination] The method according to Claim 2, characterized in that [it contains, as] component A[,] is a compound of Examples 92-119.
 8. (Amended) [Use of a combination] The method according to Claim 2, characterized in that [it contains, as] component A[,] is a compound of Examples 48 or 80.

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9. (Amended) [Medicament] A pharmaceutical composition comprising a combination of an MTP inhibitor as component A and an HMG-CoA reductase inhibitor as component B according to Claim 1 and, if appropriate, one or more further suitable components.
 10. (Amended) [Medicament] A pharmaceutical composition according to Claim 9, characterized in that it contains, as component A, the active compound 2-cyclopentyl-2-[4-(2,4-dimethyl-pyrido[2,3-b]indol-9-ylmethyl)-phenyl]-N-(2-hydroxy-1-phenyl-ethyl)-acetamide or 2-cyclopentyl-2-[4-(2,4-dimethyl-pyrimido[1,2-a]indol-10-ylmethyl)-phenyl]-N-(2-hydroxy-1-phenyl-ethyl)-acetamide and, as component B, the active compound atorvastatin, cerivastatin, simvastatin, pravastatin, lovastatin, fluvastatin, itavastatin or ZD 4522.
 11. (Amended) [Medicament] A pharmaceutical composition according to Claim 9, characterized in that it contains, as component A, the compound (2S)-2-cyclopentyl-2-[4-(2,4-dimethyl-pyrido[2,3-b]indol-9-ylmethyl)-phenyl]-N-(2-(1R)-hydroxy-1-phenyl-ethyl)-acetamide.
 12. (Amended) [Process] A process for the production of [medicaments] a pharmaceutical composition according to Claim 9, characterized in that the components A and B are converted into a suitable administration form with excipients and vehicles and, if appropriate, with further components.